

# CTG TGT GTC CTG ATG GTC CTG ATG TTG GGC AGA GTC CTG

CTC ACG ATG ACC AGG TGT GCT CTG CTG TTG CTG ATG GTC CTG ATG TTG GGC AGA GTC CTG  
 M T R C A L L L L M M V L M L G R V L  
 10  
 20 GTT GTC CCA GTG ACC CCT ATC CCA ACC TTC CAG CTG CGC CCT CAG AAT TCT CCC CAG ACC  
 V P V T P I P T F Q L R P Q N S P Q T  
 30  
 40 ACT CCC CGA CCT GCG GCC TCA GAG AGC CCC TCA GCT GCT CCC ACA TGG CCG TGG GCT GCC  
 T P R P A A S E S P S A A P T W P W A A  
 50  
 60 CAG AGC CAC TGC AGC CCC ACC CGC CAC CCT GGC TCG CGC ATT GTC CTA TCG CTG GAT GTC  
 Q S H C S P T R H P G S R I V L S L D V  
 70  
 80 CCC ATC GGC CTC TTG CAG ATC TTA CTG GAG CAA GCC CGG GCC AGG GCT GCC AGG GAG CAG  
 P I G L L Q I L L E Q A R A R A A R E Q  
 90  
 100 GCC ACC AAC GCC CGC ATC CTG GCC CGT GTC GGC CAC TGC TGA GCC TGA GAG AGG GGG  
 A T T N A R I L A R V G H C \* (SEQ ID NO:2)  
 110  
 TCA CAG TGA TAG GGC CAC CCT GGA TGG GAA GAC CTG GAG (SEQ ID NO:1)

Fig. 1

# Sequence Alignment

|     |     |     |     |     |     |     |     |     |     |     |     |     |     |               |     |     |     |     |     |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|---------------|-----|-----|-----|-----|-----|
| CTC | ACG | ATG | ACC | AGG | TGT | GCT | CTG | CTG | TTG | CTG | ATG | GTC | CTG | ATG           | TTG | GGC | AGA | GTC | CTG |
| M   | T   | R   | C   | A   | L   | L   | L   | L   | L   | M   | V   | L   | M   | L             | G   | R   | V   | L   |     |
| 20  |     |     |     |     |     |     |     |     |     |     |     |     |     |               |     |     |     |     |     |
| GTT | GTC | CCA | GTG | ACC | CCT | ATC | CCA | ACC | TTC | CAG | CTC | CGC | CCT | CAG           | AAT | TCT | CCC | CAG | ACC |
| V   | P   | V   | T   | P   | I   | P   | T   | F   | Q   | L   | R   | P   | Q   | N             | S   | P   | Q   | T   |     |
| 30  |     |     |     |     |     |     |     |     |     |     |     |     |     |               |     |     |     |     |     |
| ACT | CCC | CGA | CCT | GCG | GCC | TCA | GAG | AGC | CCC | TCA | GCT | GCT | CCC | ACA           | TGG | CCG | TGG | GCT | GCC |
| T   | P   | R   | P   | A   | A   | S   | E   | S   | P   | S   | A   | A   | P   | T             | W   | P   | W   | A   | A   |
| 40  |     |     |     |     |     |     |     |     |     |     |     |     |     |               |     |     |     |     |     |
| CAG | AGC | CAC | TGC | AGC | CCC | ACC | CGC | CAC | CCT | GGC | TCG | CGC | ATT | GTC           | CTA | TCG | CTG | GAT | GTC |
| Q   | S   | H   | C   | S   | P   | T   | R   | H   | P   | G   | S   | R   | I   | V             | L   | S   | L   | D   | V   |
| 50  |     |     |     |     |     |     |     |     |     |     |     |     |     |               |     |     |     |     |     |
| CCC | ATC | GGC | CTC | TTG | CAG | ATC | TTA | CTG | GAG | CAA | GCC | CGG | GCC | AGG           | GCT | GCC | AGG | GAG | CAG |
| P   | I   | G   | L   | L   | Q   | I   | L   | L   | E   | Q   | A   | R   | A   | R             | A   | A   | R   | E   | Q   |
| 60  |     |     |     |     |     |     |     |     |     |     |     |     |     |               |     |     |     |     |     |
| GCC | ACC | AAC | ACC | GCC | CGC | ATC | CTG | GCC | CGT | GTC | GGC | CAC | TGC | TGA           | GCC | TGA | GAG | AGG | GGG |
| A   | T   | N   | A   | R   | I   | L   | A   | R   | V   | G   | H   | C   | *   | (SEQ ID NO:2) |     |     |     |     |     |
| 70  |     |     |     |     |     |     |     |     |     |     |     |     |     |               |     |     |     |     |     |
| TCA | CAG | TGA | TAG | GGC | CAC | CCT | GGA | TGG | GAA | GAC | CTG | GAG |     |               |     |     |     |     |     |
| 80  |     |     |     |     |     |     |     |     |     |     |     |     |     |               |     |     |     |     |     |
| 100 |     |     |     |     |     |     |     |     |     |     |     |     |     |               |     |     |     |     |     |

Fig. 2



# Fig. 4A

MTRWALVVFVVLMLDRILFVPGTPIPTFFQLLPQNSLETP 40  
 SSVTSESSSGTTTGPSASWSNSKASPYLDTRVILSLDVPI 80  
 GLLRILLEQARYKKAARNQAATNAQILAHVGRR (SEQ ID NO: 10) 112

Fig. 4A

|               |   |   |                 |
|---------------|---|---|-----------------|
| mouse Ucn II: | VILSLDVPTIGLLRITLLEQARYKKAARNQAATNAQILAHV | • | (SEQ ID NO: 10) |
| human URP:    | IVLSLDVPTIGLLITLLEQARARAAREQATTNARILARV   | • | (SEQ ID NO: 11) |
| fish URP:     | LTLSLDVPTNIMNVIFDVAKKNILRAKAAENARILAHV    | • | (SEQ ID NO: 12) |
| rat Ucn:      | DDPPLSILTFHLLRTLLEILARTQSQRERAEQNRIIFDSV  | • | (SEQ ID NO: 13) |
| r/hCRF:       | SEEPPISLDLTFHLLREVLLEMAAEQLAQQAHSNRKLMETI | • | (SEQ ID NO: 14) |

Fig. 4B

Displacement of  
 $^{125}$ I-Sauvagine binding to  
CRF-R1

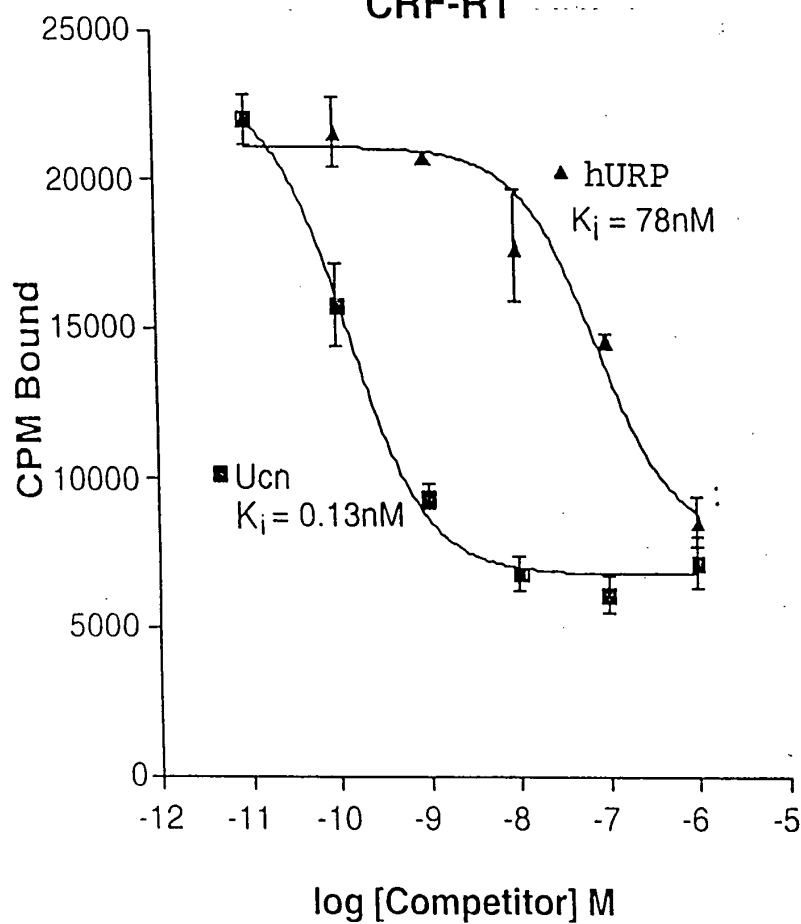


Fig. 5A

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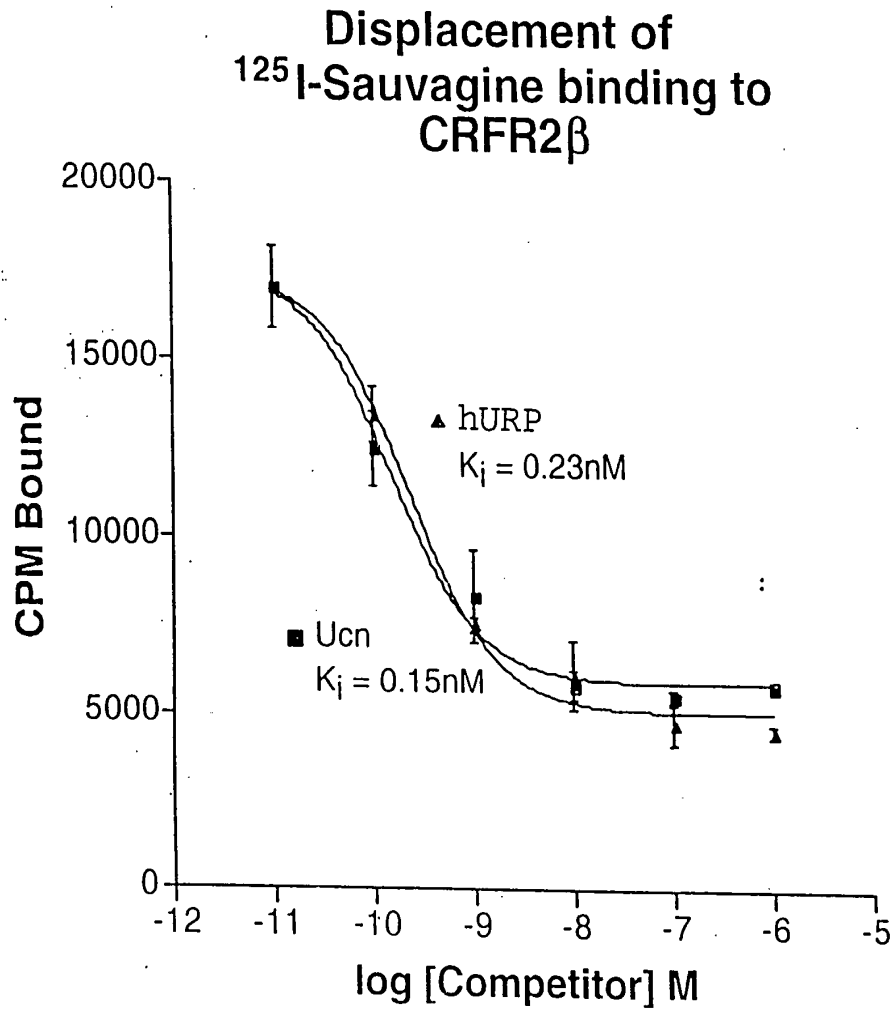


Fig. 5B

**A**

mip

pm

v3

**B**

CBL

v4

LC

**C**

VII

men

**Fig. 6**

**Fig. 7**



This electron micrograph shows a presynaptic terminal, labeled 'VII', which is a dense, electron-lucent structure. Below it, the postsynaptic membrane is visible, labeled 'men'. The image is characterized by a high level of contrast and a grainy texture, typical of electron microscopy.

**Fig. 8**

00040407404

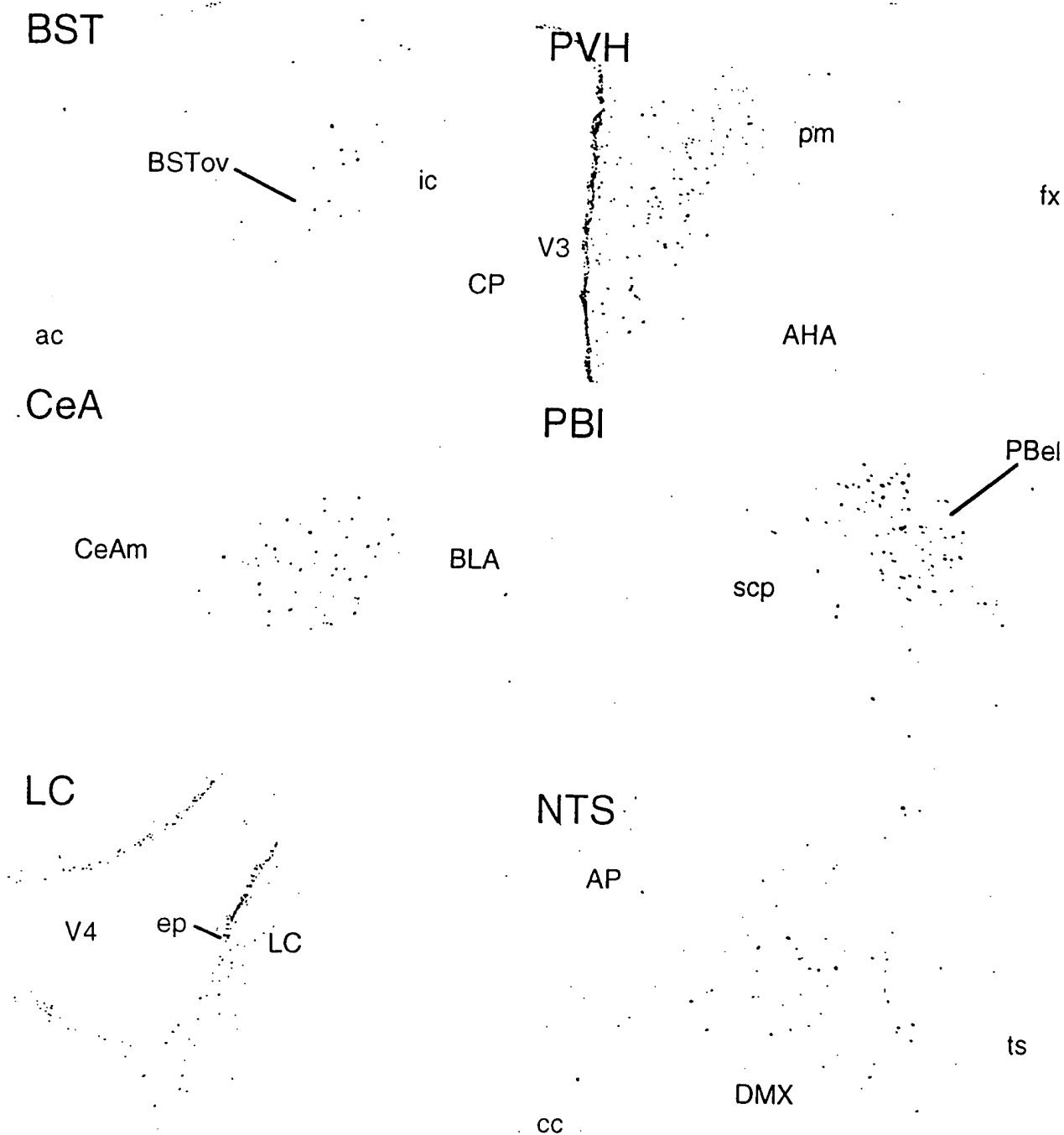


Fig. 9

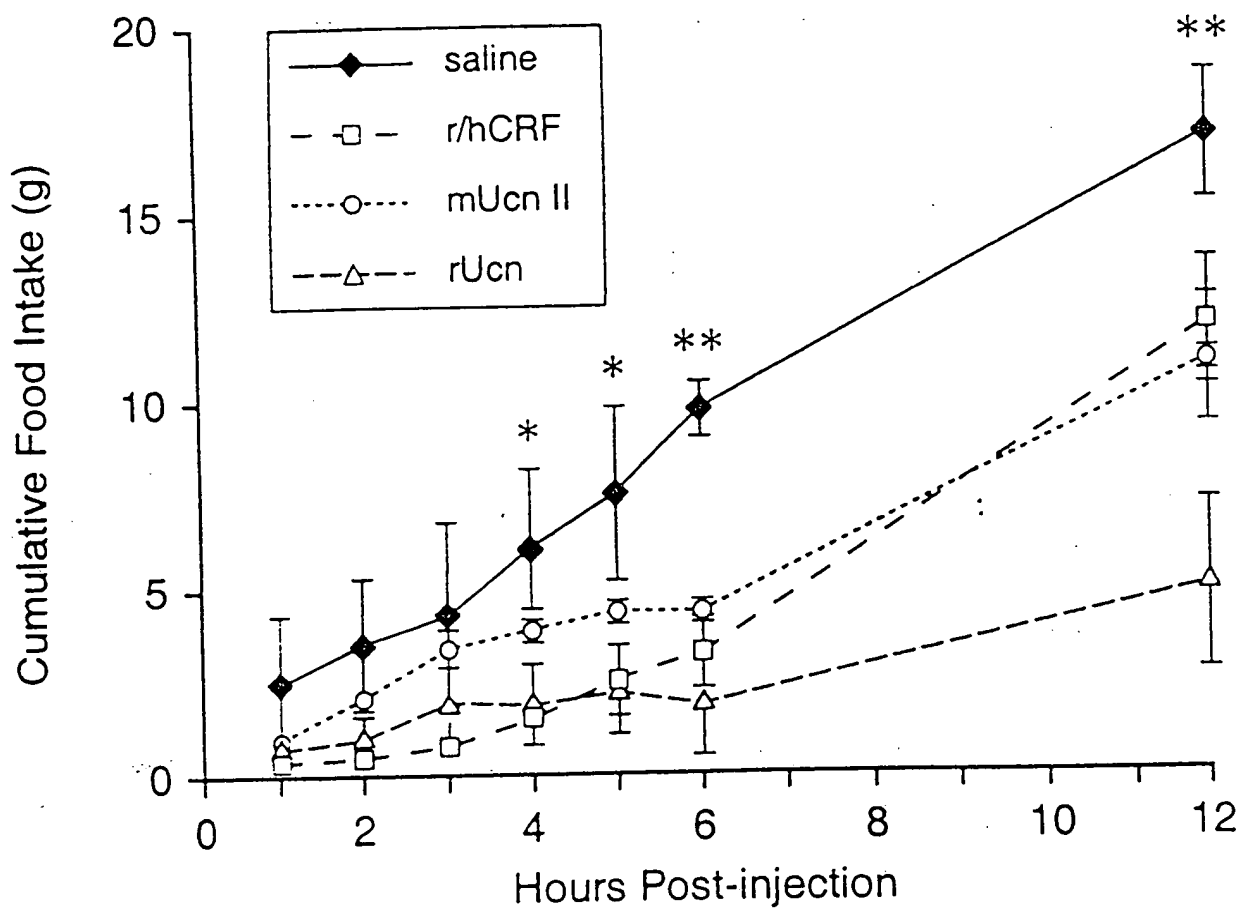


Fig. 10A

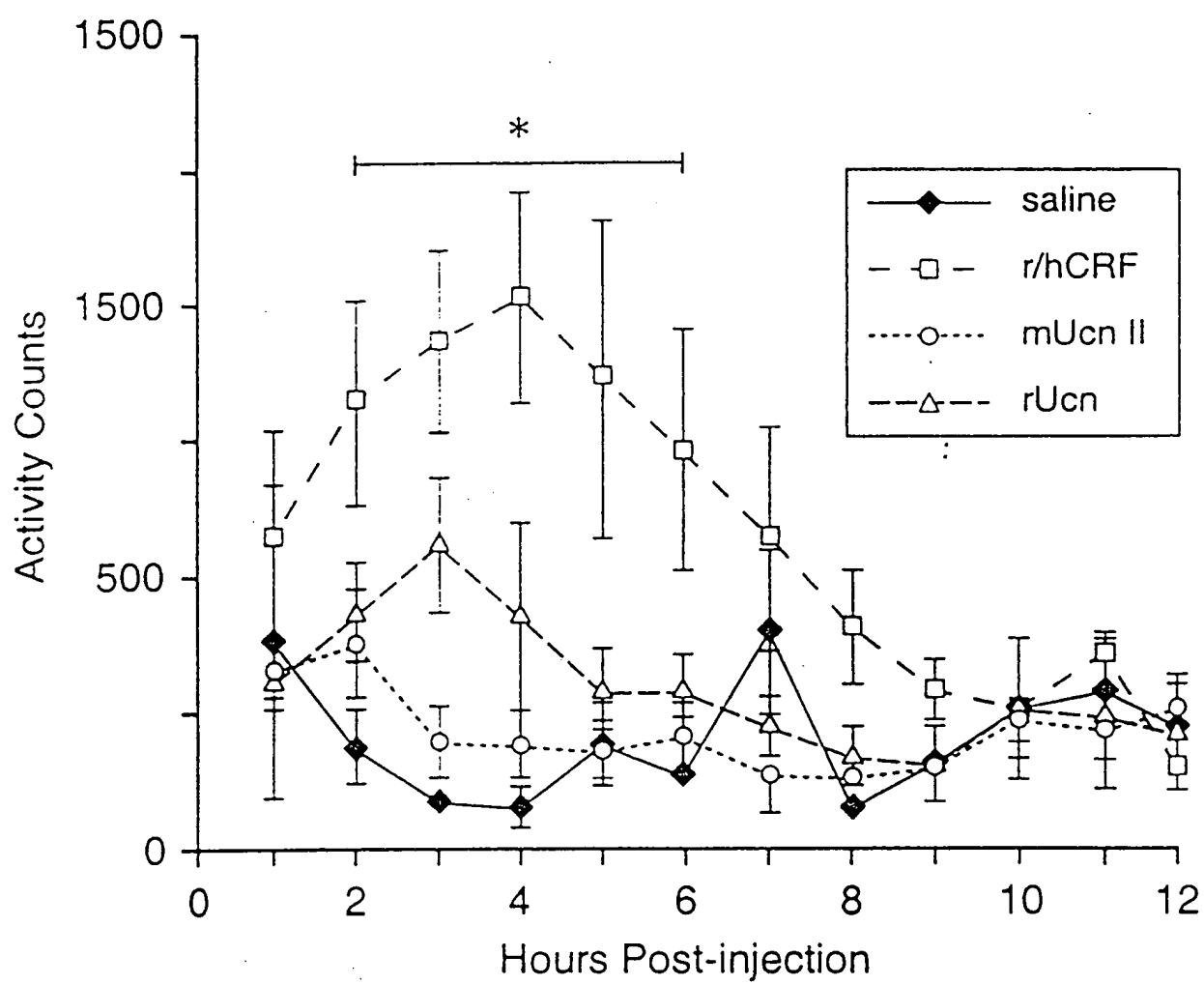


Fig. 10B

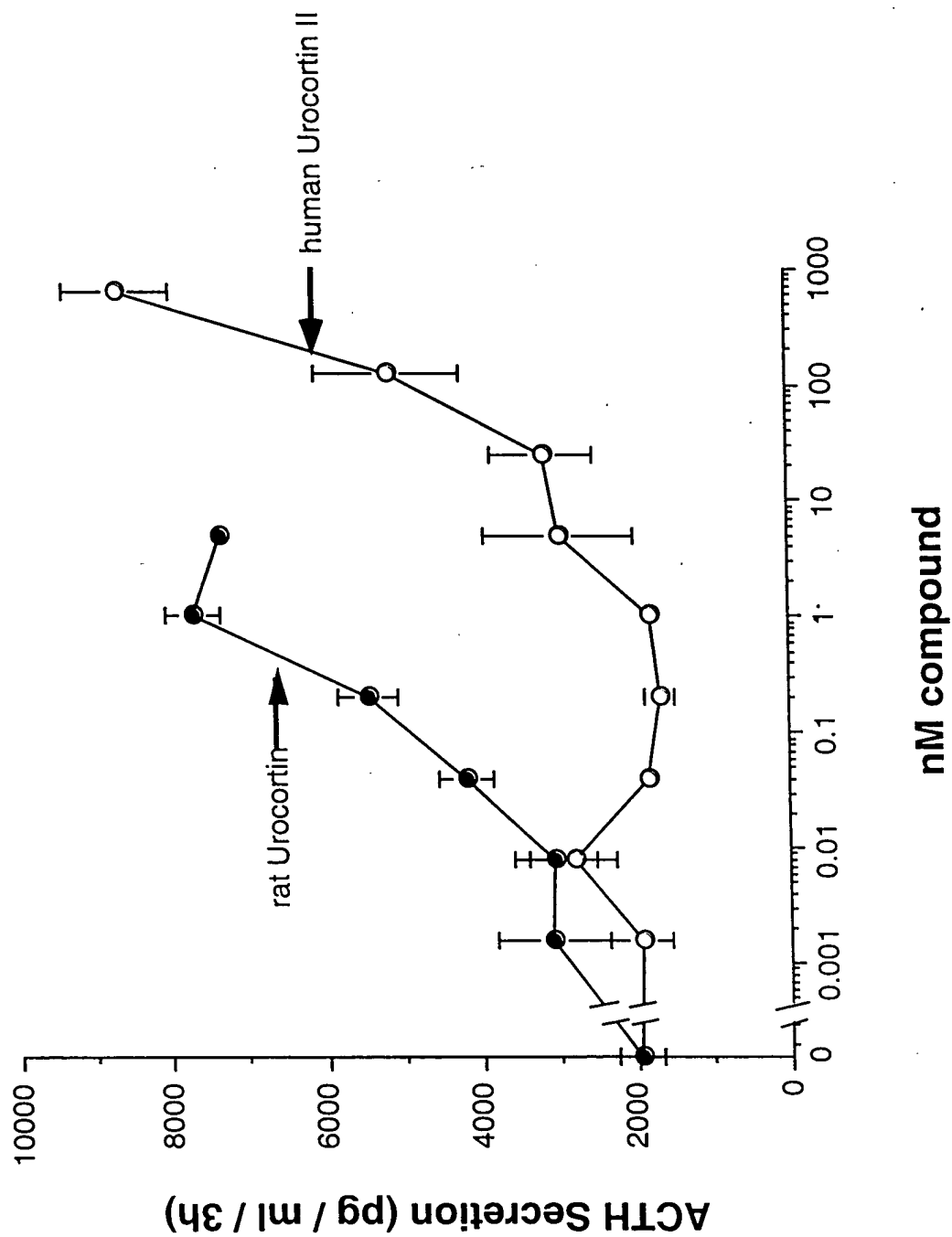


Fig. 11

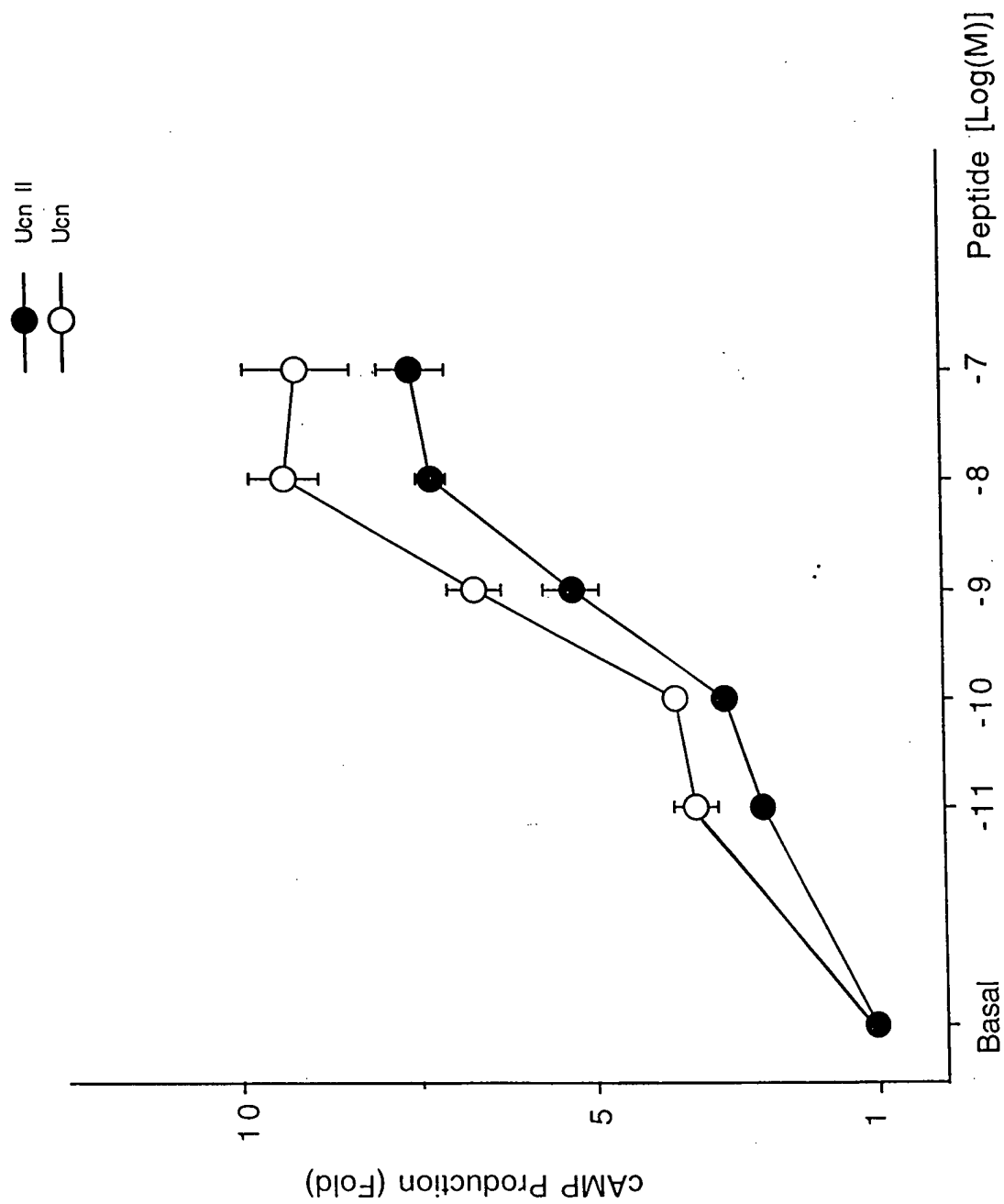
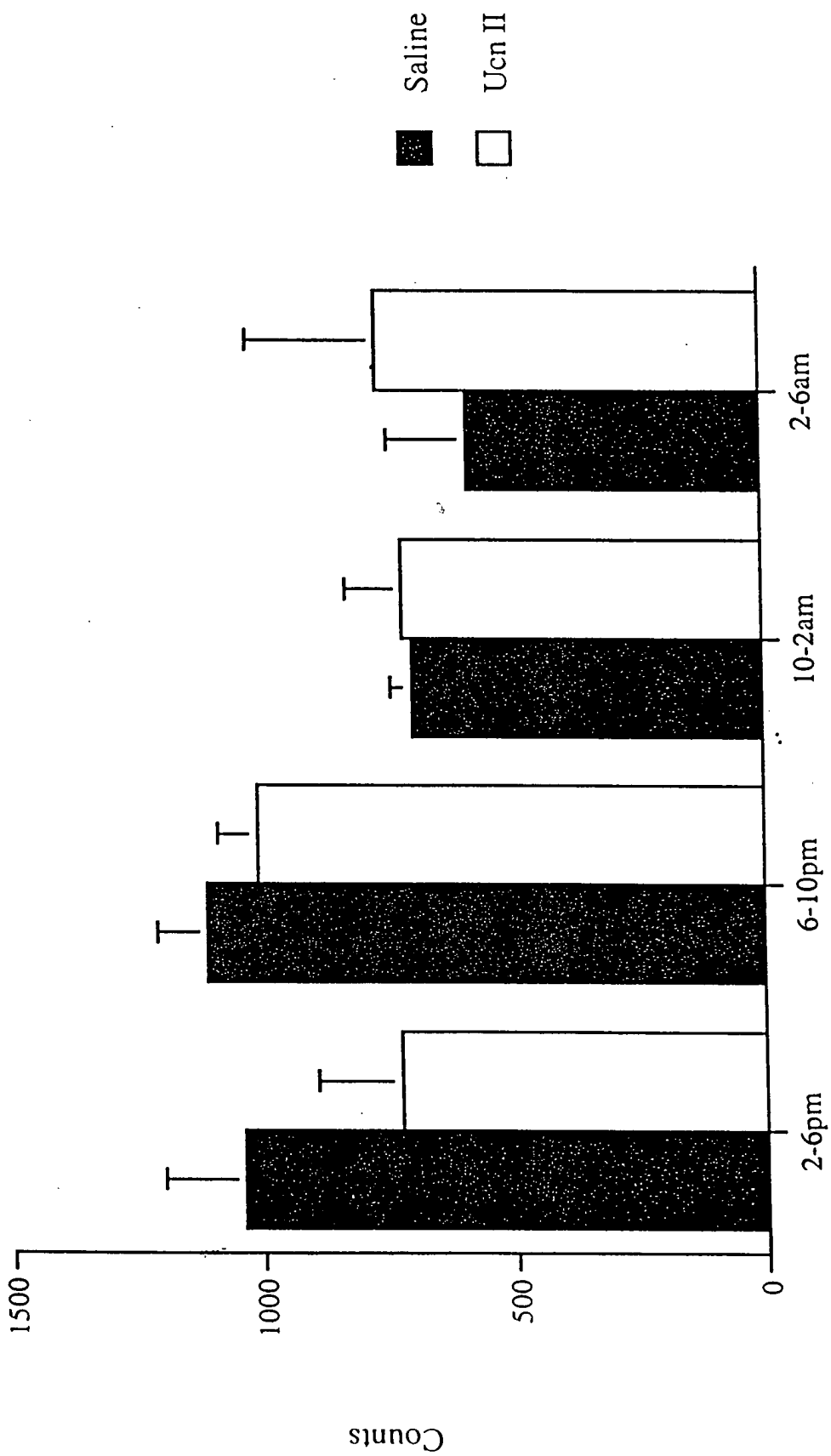
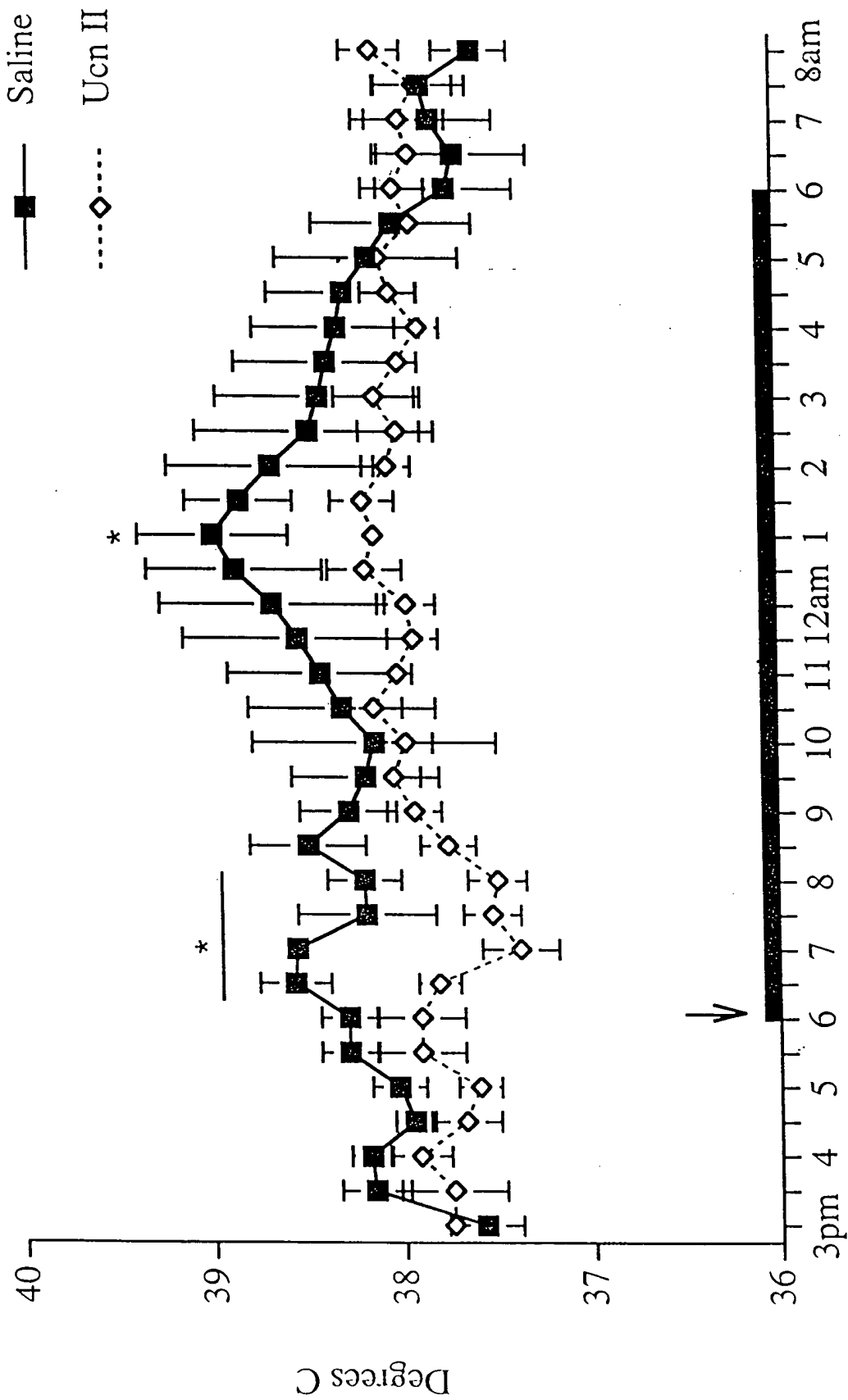


Fig. 12



Time of Day

Fig. 13



Time of Day

**Fig. 14**



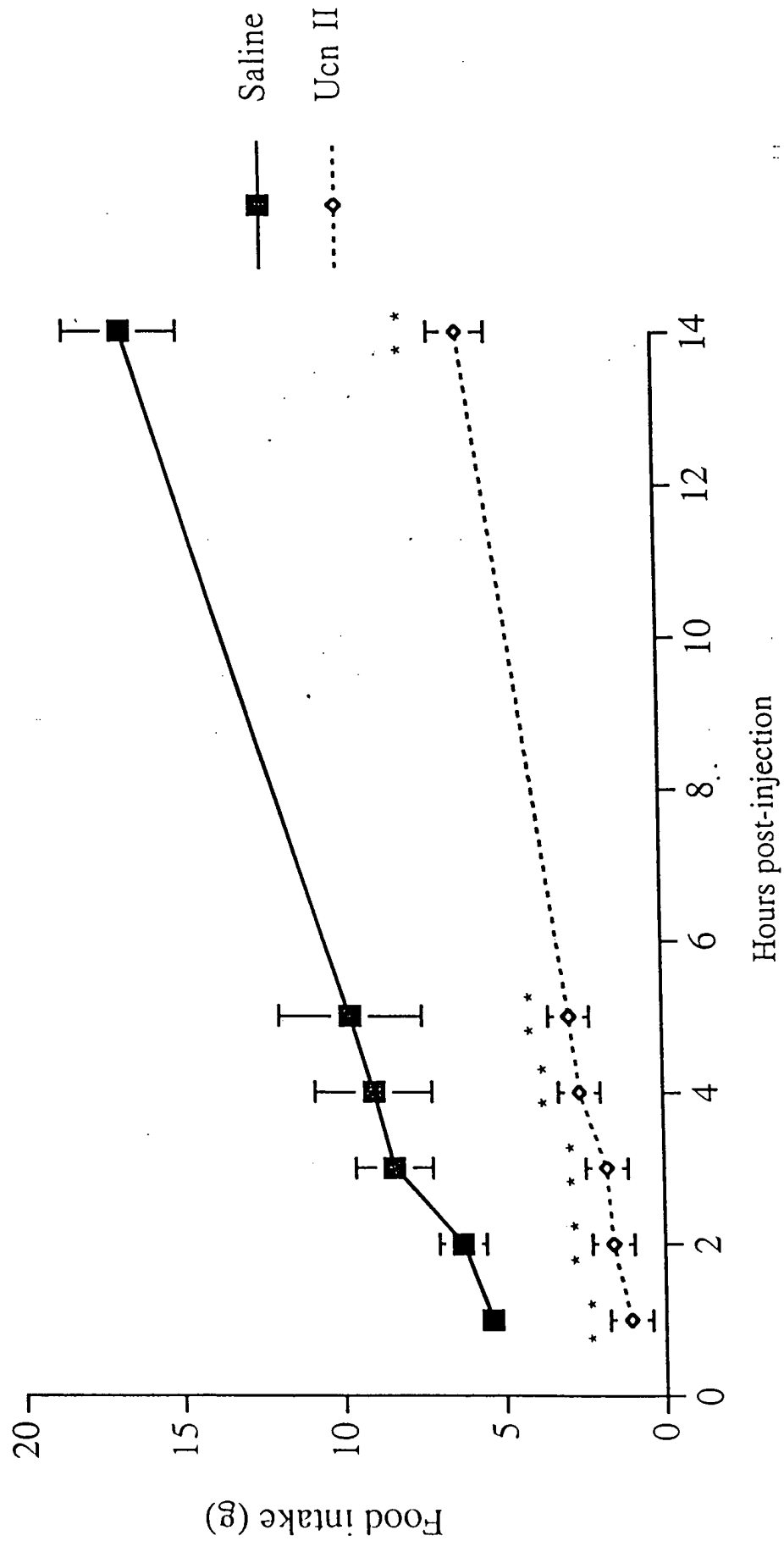
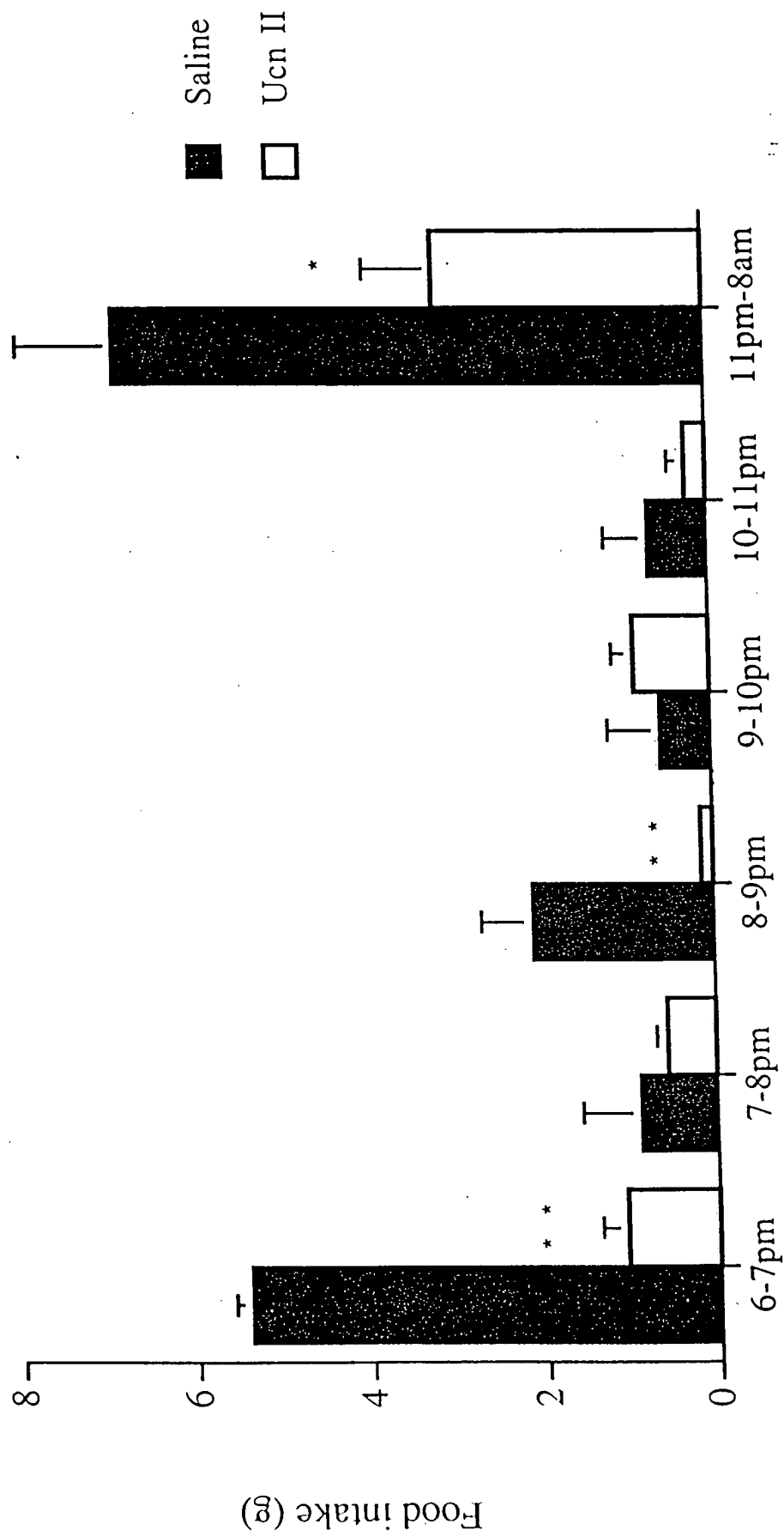


Fig. 15A

Hours post-injection



Time of day

Fig. 15B

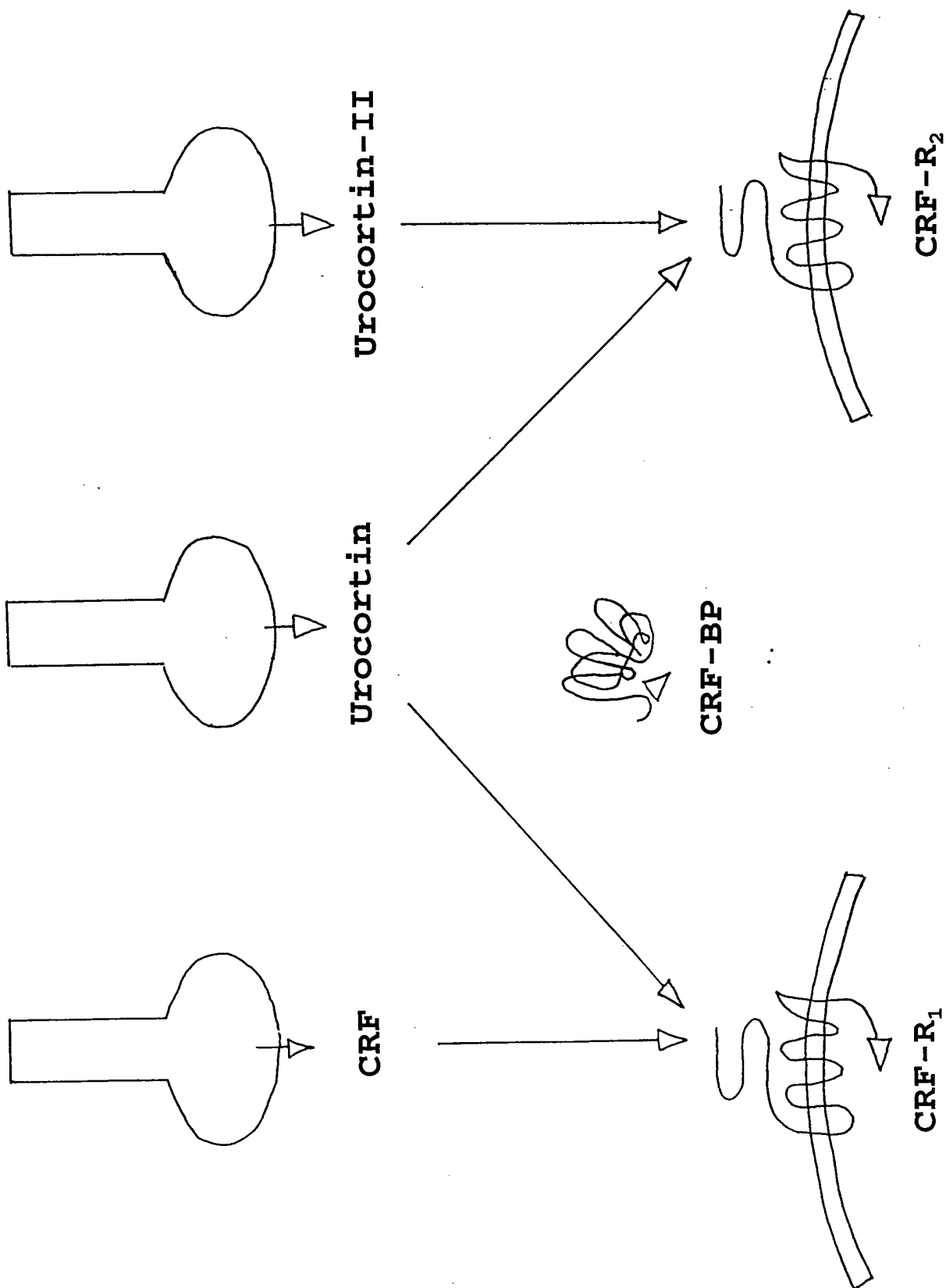


Fig. 16